IN THE CLAIMS

 (Currently Amended) A method of synthesis of a chemical compound having the formula A-B-C

where the A is a chemiluminescent moiety comprising a phthalhydrazide,

B is an energy acceptor moiety, and

C is a biologically active moiety

comprising the steps of

forming a benzophenone,

forming a diaryl ethylene, and

performing comprising the steps of at least one of

- (a) forming benzophenone;
- (b) forming a diaryl ethylene:
- (c) attaching a precursor to generate a phthalhydrazide; such as phthalimide, aminophthalic acid diester, aminophthalic acid dihydrazide, aminophthalic anhydride and phthalhydrazide protected by a hydrolyzable group to form the precursorethylene conjugate, and
- (d) condensing two ethylene-precursor conjugates to form a precursor-pentadiene conjugate; -and
- (e) (b) condensing two diaryl ethylene to form a pentadiene: and
- (f) attaching a precursor to a <u>pentadiene</u> generate a phthalhydrazide such as phthalimide, aminophthalic acid diester, aminophthalic acid dihydrazide, aminophthalic anhydride and phthalhydrazide protected by a hydrolyzable group, to form a the precursor-pentadiene conjugate;; and
- (g) converting the <u>a</u> precursor to the phthalhydrazide by at least one of the corresponding reactions

phthalimide with hydrazine,

aminophthalic acid diester with hydrazine,

aminophthalic anhydride with hydrazine, and

hydrolysis of phthalhydrazide protected by a hydrolyzable group to form a carrier compound, and

reacting the carrier compound with the biologically active moiety to form a corresponding conjugate.

2. (Original) The method of synthesis of the compound of claim 1 wherein the compound

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serves to delivery the C moiety to a desired biological compartment.

- (Original) The method of synthesis of the compound of claim 1 wherein the compound is a prodrug.
- 4. (Original) The method of synthesis of the compound of claim 3 wherein the compound serves as a prodrug for at least one of antiviral agents for the treatment of viral infections and anticancer agents for the treatment of cancers.
- 5. (Original) The method of synthesis of the compound of claim 4 wherein the compound serves as a prodrug for the treatment of at least one of the group of viruses comprising Human Immunodeficiency Virus (HIV), herpes viruses such as Herpes Simplex Virus, (HSV), Epstein-Barr Virus (EBV), Varicella Zoster (VZV), Cytomegalovirus (CMV), HSV-6, and HSV-8 (Kaposi's sarcoma), Human Papilloma Virus (HPV), rhinoviruses, and hepatitis-linked viruses.
- (Original) The method of synthesis of the compound of claim 4 wherein the compound serves as a prodrug for the treatment of at least one of the group of cancers comprising colon, breast, lung, renal, retinal, and skin.
- (Original) The method of synthesis of the compound of claim 3 wherein the prodrugs have increased bioavailability.
- (Original) The method of synthesis of the compound of claim 2 wherein the compound is a cellular permeant prodrug.
- (Original) The method of synthesis of the compound of claim 8 wherein intracellular drug release occurs when the prodrug reacts with cellular free radicals via a mechanism involving chemiluminescence, photochromism, and intramolecular energy transfer.
- (Original) The method of synthesis of the compound of claim 1 wherein the C moiety is a pharmaceutical agent or drug.
- 11. (Original) The method of synthesis of the compound of claim 10 wherein the pharmaceutical agent is at least one of the group of antilipidemic drugs, anticholesterol drugs, contraceptive agents, anticoagulants, anti-inflamatory agents, immuno-suppressive drugs,

antiarrhythmic agents, antineoplastic drugs, antihypertensive drugs, epinephrine blocking agents, cardiac inotropic drugs, antidepressant drugs, diuretics, antifungal agents, antibacterial drugs, anxiolytic agents, sedatives, muscle relaxants, anticonvulsants, agents for the treatment of ulcer disease, agents for the treatment of asthma and hypersensitivity reactions, antithroboembolic agents, agents for the treatment of muscular dystrophy, agents to effect a therapeutic abortion, agents for the treatment of anemia, agents to improve allograft survival, agents for the treatment of disorders of purine metabolism, agents for the treatment of ischemic heart disease, agents for the treatment of opiate withdrawal, agents which activate the effects of secondary messenger inositol triphosphate, agents to block spinal reflexes, and antiviral agents including a drug for the treatment of AIDS.

- 12. (Original) The method of synthesis of the compound of claim 1 wherein the C moiety is released by an oxidation reduction reaction with the target cell's electron carriers or by reaction with free radicals produced as a consequence of electron transport.
- 13. (Original) The method of synthesis of the compound of claim 12 wherein the C moiety is released into a desired compartment in active form.
- 14. (Original) The method of synthesis of the compound of claim 13 wherein the released C mojety has a greater therapeutic effect or therapeutic ratio relative to the free C agent alone.
- 15. (Original) The method of synthesis of the compound of claim 14 wherein the released C moiety has a greater therapeutic effect or therapeutic ratio relative to the free C agent alone as a consequence of at least one of altered pharmacokinetics or pharmacodynamics such as a desirable kinetics of release, a resistance to inactivation or excretion, greater solubility, enhanced absorption, a diminished toxicity, or greater access to the cellular or biological compartment which is the site of action of C.
- 16. (Original) The method of synthesis of the compound of claim 1 wherein A represents a functionality which undergoes at least one of

an oxidation reduction reaction where electrons are transferred directly between A and the target cell's electron carriers, and

a reaction with free radicals of oxygen which are produced as a consequence of electron transport

such that an excited state is produced in A as a consequence of its participation in one of

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these reactions.

- 17. (Original) The method of synthesis of the compound of claim 16 wherein A undergoes intramolecular energy transfer from its own excited state to the B functionality which is an energy acceptor.
- 18. (Original) The method of synthesis of the compound of claim 17 wherein upon receiving energy from A, B achieves an excited state which relaxes through heterolytic cleavage of the covalent bond of B with C where C is a drug moiety which is released into the environment.
- 19. (Original) The method of synthesis of the compound of claim 18 wherein the released drug molecule effects a therapeutic functional change by a mechanism which comprises receptor mediated mechanisms including reversible and irreversible competitive agonism or antagonism including a molecule known as a suicide substrate or a transition state analogue mechanism or a noncompetitive or uncompetitive agonism or antagonism or the action is by a nonreceptor mediated mechanism including a "counterfeit incorporation-mechanism".
- 20. (Currently Amended) The A method of synthesis of a chemical the compound of claim-1 having the formula A-B-C

where wherein the A is a chemiluminescent moiety;

B is an energy acceptor moiety, and

C is a biologically active moiety

the method comprising the steps of condensing A and B to form a conjugate A-B and reacting the conjugate A-B wth C wherein the chemiluminescent moiety comprises a molecule selected from comprises at least one of the group consisting of

molecules undergoing reaction involving peroxides and oxygen free radicals,

molecules undergoing reaction involving oxidation or reduction, and

molecules undergoing both reaction with peroxides and oxygen free radicals followed by an oxidation or reduction reaction.

21. (Original) The method of synthesis of the compound of claim 20 wherein the chemiluminescent molecule comprises at least one of the group of luminol and its derivatives, lucigenin and its derivatives, Lophine and its derivatives, acridinium esters and acridans, tetraphenylpyrrole, phthalhydrazides, acyloins, biacridinium salts, vinylearbonyls, vinylnitriles, tetrakis (dimethylamino) ethylene, acyloproxides, indoles, tetracarbazoles and active oxalates.

- 22. (Original) The method of synthesis of the compound of claim 20 wherein the chemiluminescent molecule comprises at least one of the group of ruthenium chelates 2, 6-diaminopyrene, or cation radicals and molecules which follow a Chemically Initiated Electron Exchange Luminescence mechanism such as certain dioxetans and dioxetanones.
- 23. (Original) The method of synthesis of the compound of claim 20 wherein the chemiluminescent molecule comprises at least one of the group of dioxene derivatives and other compounds that form a dioxetan by reaction with superoxide and then produce efficient chemiluminescence by a CIEEL mechanism.
- 24. (Curently Amended) The method of synthesis of the compound of claim 20 wherein the chemiluminescent molecule comprises at least one of the group of

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Dioxene

Imidazole derivatives

$$R_1 \stackrel{N}{\longrightarrow} R_2$$
 R_3

Sulfonyloxamides

Indole derivatives

Tetrakis(dialkylamino)ethylene

2,5,7,8-tetraexabicyclo-[4.2.0.] octane

$$O$$
 R_2
 R_1
 O
 O

Dioxetan

Lucigenin

- 25. (Currently Amended) The method of synthesis of the compound of claim + 20 wherein the energy acceptor moiety B moiety is a photochromic compound.
- 26. (Original) The method of synthesis of the compound of claim 25 wherein the

photochromic compound comprises one which demonstrate photochromic behavior with electromagnetic radiation and bleaching agents.

- 27. (Original) The method of synthesis of the compound of claim 26 wherein the A functionality is chemiluminescent, and the B functionality is such that the photodissociative drug release spectrum of B overlaps the chemiluminescence spectrum of A.
- 28. (Original) The method of synthesis of the compound of claim 25 wherein the photochromic compound comprises a cationic dye.
- 29. (Original) The method of synthesis of the compound of claim 28 wherein the cationic dye comprises at least one of a di and triarylmethane dyes, triarylmethane lactones and cyclic ether dyes, cationic indoles, pyronines, phthaleins, oxazines, thiazines, acridines, phenazines, and anthocyanidins, and cationic polymethine dyes and azo and diazopolymethines, styryls, cyanines, hemicyanines, dialkylaminopolyenes, and other related dyes.
- 30. (Currently Amended) The method of synthesis of the compound of claim 28 wherein the cationic dye comprises at least one of

Malachite Green	42000
Helvetia Green	42020
Basic Blue 1	42025
Brilliant Blue	
Setoglaucine	
Basic Green 1	42040
Brilliant Green	
Acid Blue 1	42045
Xylene Blue VS	
Patent Blue V	
Alphazurine 2G	
Acid Blue 3	42051
Brilliant Blue V	
Patent Blue V	
Food Green 3	42053

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FDC Green 3	
Acid Green 6	42075
Light Green SF Bluish	
Acid Blue 7	42080
Xylene Blue AS	
Patent Blue A	
Acid Green 3	42085
Acid Blue 9	42090
Erioglaucine	
Acid Green 5	42095
Light Green SF Yellowish	
Acid Green 9	42100
Erioviridene B	
Acid Blue 147	42135
Xylene Cyanol FF	
Basic Red 9	42500
Pararosaniline	
Basic Violet 14	42510
Fuchsin	
Magenta	
Basic Fuchsin	42510B
Basic Violet 2	42520
New Magenta	
Hoffman Violet	42530
Iodine Violet	
Basic Violet 1	42535
Methyl Violet	
Basic Violet 13	42536
Methyl Violet 6B	
Basic Violet 3	42555
Crystal Violet	
Gentian Violet	
Iodine Green	42556
Basic Blue 8	42563
Victoria Blue 4R	

Acid Blue 13	42571
Fast Acid Violet 10B	
Acid Blue 75	42576
Eriocyanine A	
Methyl Green	42585
Ethyl Green	42590
Basic Violet 4	42600
Ethyl Violet	
Acid Violet 49	42640
Wool Violet 5BN	
Acid Blue 15	42645
Brilliant Milling Blue B	
Acid Violet 17	42650
Acid Violet 6B	
Wood Violet 4BN	
Formyl Violet	
Acid Violet 5BS Conc.	
Acid Violet 19	42685
Acid Fuchsin	
Red Violet 5R	42690
Acid Blue 22	42755
Aniline Blue	
Soluble Blue	
Solvent Blue 3	42775
Solvent Blue 3	42780
Methyl Blue	
Aurin	43800
Mordant Blue 3	43820
Eriochrome Cyanine R	
Acid Green 16	44025
Naphthalene Green V	
Pontacyl Green NV Extra	
Basic Blue 11	44040
Victoria Blue R	
Basic Blue 15	44085

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Night Blue

Acid Green 50

Wool Green S

Kiton Green S. Conc.

Basic Green 3

Sevron Green B

Brilliant Blue F & R Extra

Brilliant Green Sulfonate

Hexakis (hydroxyethyl)

New Green

$$(CH_2)_2N$$
 C^* OCH_3

Phenolphthalein

Malachite Green Ethiodide

Hydroxyalkylated Pararosanilines

44090

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Hydroxyalkylated New Fuchsins

$$(CH_1)_2N$$
 — $C^*(C_6H_5)_2$

Doebner's Violet

$$\left(H_2 N - \left(\begin{array}{c} \\ \\ \\ \end{array} \right)_2 C^* C_6 H_5 \right.$$

New Red

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Bis(hydroxyethyl) Doebner's Violet

$$- \left\{ \text{HOCH}_2\text{CH}_2\text{NH} - \left\{ \begin{array}{c} \\ \\ \end{array} \right\}_2 \text{C*C}_0\text{H}_5$$

"New Magenta"

Tetrakis(hydroxyethyl) Doebner's Violet

Trichloro Crystal Violet

$$(CH_3)_2N$$
 C^*

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Slow Red

$$(CH_0)_{1}N \longrightarrow C^2 \longrightarrow CCH_1$$

$$C_{1}H_1NH \longrightarrow J_3$$

$$(C_{2}H_2)_{2}N \longrightarrow J_2$$

$$(C_{3}H_3)_{2}N \longrightarrow J_2$$

$$(C_{3}H_3)_{2}N \longrightarrow J_3$$

$$(C_{3}H_3)_{2}N \longrightarrow J_3$$

$$(C_{3}H_3)_{2}N \longrightarrow J_3$$

$$(C_{3}H_3)_{3}N \longrightarrow J_3$$

$$(C_{3}H_3)_{3}N \longrightarrow J_3$$

$$(C_{3}H_3)_{3}N \longrightarrow J_3$$

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$$\begin{pmatrix} (\operatorname{CH}_3)_2 N & & & \\ & \ddots & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & &$$

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^a Only the cyanide, bisulfite, and hydroxide ions are considered, regardless of the other anions present in the solution.

b More detailed descriptions of the compositions of photochromic materials tested are given in Macnair's review [255; tables 1A-4].

c Ethanol.

d Diethyl ether.

c 1,2-Dichloroethane.

f 1,1-Dichloroethane, cyclohexane-1,1-dichloroethane, or cyclohexane-1,2-dichloroethane mixtures.

g Benzene.

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- h Dimethylsulfoxide, neat and aqueous.
- i Acetone.
- JACETIC ACID.
- k Ethyl acetate.
- 1 Ethyl bromide.
- m 2-Methoxyethanol.
- n Chloroform
- O Ethanol with KCN.
- p Ethanol with KOH.
- ^q Carboxylic acids-acetic to stearic; hydrocinnamic acid; ethyl and butyl acid phthalates.
- Getadecylnitrile, tributyl phosphate, aniline, 2-(p-tert-butylpheno xy)ethanol, tetraethyleneglycol dimethyl ether, or poly(ethylene glycols).
- S Amides-formamide to stearamide; methylformamide or methylacetamide; dimethyl- or diethyl-formamide or acetamide.
- ¹ Three-to-one solutions of cellulose acetate with any of the following five-to-one plasticizer mixtures: butyl stearate, Polyethylene Glycol 600-butyl acetoxystearate, butyl stearate, or Dowanol EP-butyl acetoxystearate.
- " Water containing SO2
- v Water containing bisulfite and papain.
- w Poly(vinyl alcohol) with dimethylsulfoxide (5:1).
- x Films, containing residual solvent, cast from the following solutions: ethanol-acetone solutions of vinyl acetate-vinyl alcohol copolymer; aqueous poly(vinyl alcohol); aqueous poly(vinyl pyrrolidone); or aqueous methyl vinylether-maleic acid copolymer.
- y Methanol-dioxane with aqueous NH4 HSO3.
- z Paper impregnated with a toluene solution of poly(methyl methacrylate), stearic acid, and 2-(p-tert-butylphenoxy)ethanol, then dried.
- aa Intramicellar impregnation of cellulose with the following swelling agents: n-propylamine, n-butylamine, n-hexylamine, 2-aminoethanol, dimethylformamide, acetic acid, dimethylsulfoxide, methylacetamide, dimethylacetamide, or formamide.
- bb Films cast from an approximately 4:3 mixture of a 20% solution and cellulose acetate butyrate in toluene-ethyl acetate (1:1) and triallycyanurate in dioxane.
- CC FILMS CAST FROM A 2:1 MIXTURE OF A 25% SOLUTION OF CELLULOSE ACETATE BUTYRATE IN TOLUENE-ETHYL ACETATE (1:1) AND THE TITANIUM ESTERS OF N.N.N', N'-TETRAKIS(2-HYDROXYPROPYL) ETHYLENEDIAMINE.
- ^{dd} Pure water

- ee Films cast from aqueous gelatin or other hydrocolloids.
- ff Dimethylsulfoxide with methanolic KCN.
- gg 2-Methoxyethanol with methanolic KCN.
- hh Water or aqueous methanol containing bisulfite.
- ii Paper impregnated with m-dimethoxybenzene, acetonitrile, acetic acid, or phenyl methyl carbinol.
- jj Ethanol-benzene.
- kk Aqueous ethanol, methanol, aqueous methanol, aqueous acetone, benzene-methanol, carbon tetrachloride-methanol, cyclohexane-methanol, or chloroform-methanol.
- ^{II} Films cast from 3:1 solutions of cellulose acetate and either Polyethylene Glycol 600 .RTM. or ethylene glycol phenyl ether as plasticizer.
- mm Films, containing residual solvent, cast from solutions of either cellulose acetate in 2-methoxyethanol or poly(vinyl alcohol) in aqueous ethanol.
- ⁿⁿ Films, containing residual solvent, cast from solutions of either cellulose acetate butyrate in 2-methoxyethanol or poly(vinyl acetate) in methanol.
 - ⁰⁰ Ethanol containing ammonia.
 - pp Aqueous methanol containing NH4 HSO3 and urease.
 - ^{qq} Aqueous methanol containing NH₄ HSO₃, with or without sodium dithionite.
 - " Aqueous acid at pH 1.
 - ss Aqueous ammonia containing KCN.
 - $^{\mathfrak{tt}}$ Paper impregnated with aqueous solutions with or without hydrocolloids.
 - uu 2-Methoxyethanol containing HCl.
 - vv Aqueous methanol containing NH4 HSO3, and glucose oxidase.
 - ww 9:1 Methanol-water.
 - xx Aqueous NaOH.]]

$$(CH_{3})_{2}N = (CH_{3})_{2}N = (CH_{3})_{2}$$

175

Photochromic Polymethine Dyes

$$(CH_3)_2N \xrightarrow{C_1} CH = CH)_n - CH = C \xrightarrow{I} N(CH_3)_2$$

Ar	n	
C ₆ H ₅	0, 1, 2	
4-(CH ₃) ₂ NC ₆ H ₄	0, 1, 2	
4-(CH ₃) ₂ CHC ₆ H ₄	0, 1, 2, 3, 4	
4-CH ₃ OC ₆ H ₄	0, 1, 2	
4-C ₄ H ₉ OC ₆ H ₄	0, 1, 2	
3-CH ₃ C ₆ H ₄	1, 2	
4-t-C ₄ H ₉ C ₆ H ₄	1, 2	
4-C ₂ H ₅ OC ₆ H ₄	1, 2	
4-C ₅ H ₁₁ C ₆ H ₄	1, 2	
4-FC ₆ H ₄	1	
4-Fsub ₃ CC ₆ H ₄	1	
2-(C ₆ H ₅)2NC6H4	1	
3,4-H ₂ N(OCH ₃)C ₆ H ₃	1	
2-Naphthyl	1, 2	
4-ClC ₆ H ₄	2	
2,4-Cl ₂ C ₆ H ₃	2	
1-Naphthyl	2	

$$\alpha$$
, α -bis(p-dimethylaminophenyl)polyenes
$$(CH_3)_2N \xrightarrow{\qquad \qquad \qquad \qquad } C^* \xrightarrow{\qquad \qquad } N(CH_3)_2$$

R

$$-CH = C - N(CH_0)_2$$

$$-CH = CH - N(CH_0)_2$$

$$-CH = CH - N(CH_0)_2$$

$$-CH = CCH_{2} \longrightarrow N(CH_{2}CH_{2}C)_{2}$$

$$-CH = CH \longrightarrow N(C_{2}H_{2})_{2}$$

$$-CH = CH \longrightarrow N(C_{2}H_{2})_{2}$$

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Miscellaneous polyenes

Basic Violet 7

Basic Red 13

Basic Red 14 Basic Red 15 Basic Violet 15

$$(CH_{3})N \qquad N(CH_{3})2$$

$$C = CH - CH = CH - C$$

$$(CH_{3})N \qquad N(CH_{3})2$$

$$(CH_{i})_{2N} = CH - CH = CH - CH_{i}$$

$$(CICH_{i}CH_{i})_{2N} = CH_{i}$$

(CH₃)₂N

(CH₃)₂N

$$(CH_{i})_{i}N$$

$$CH=CH-CH$$

$$CIO_{i}$$

$$(CH_{i})_{i}N$$

$$C=CH=CH$$

$$CIO_{i}$$

$$CIO_{i}$$

$$CIO_{i}$$

$$CIO_{i}$$

$$CIO_{i}$$

$$CIO_{i}$$

$$CIO_{i}$$

$$CH_{i})_{i}N$$

$$C=CH-CH$$

$$CIO_{i}$$

$$CH_{i})_{i}N$$

$$CIO_{i}$$

$$CH_{i})_{i}N$$

$$CH_{i}N$$

$$C$$

N(CH₃)₂

$$(CH_{0})N$$

$$C = CH - N = N$$

$$CH_{0} = N$$

$$CH_{0} = N$$

$$CH_{0} = N$$

$$CH_{0} = N$$

$$N(CH_{0})_{2}$$

$$N(CH_{0})_{2}$$

$$N(CH_{0})_{2}$$

$$N(CH_{0})_{2}$$

$$N(CH_{0})_{2}$$

$$N(CH_{0})_{2}$$

$$N(CH_{0})_{2}$$

$$N(CH_{0})_{3}$$

$$N(CH_{0})_{4}$$

$$\begin{array}{c|c} H_1C & CH_3 \\ \hline H_1N & C & \\ \hline \\ NiH_2 & CH_3 \\ \hline \\ NiH_2 & CH_3 \\ \hline \\ NiCH_3)_2 & CI \\ \hline \\ NiCH_3)_3 & CI \\ \hline \\ NiCH_3)_4 & CI \\ \hline \\ NiCH_3)_3 & CI \\ \hline \\ NiCH_3)_4 & CI \\ \hline \\ NiCH_3)_4 & CI \\ \hline \\ NiCH_3)_5 & C$$

$$N_{i}C_{i}H_{i})CH_{i} - \sum_{N_{i}C_{i}H_{i}}N_{i}C_{i}H_{i}CH_{i} - \sum_{N_{i}C_{i}H_{i}}N_{i}H_{i} - \sum_{N_{i}C_{i}}N_{i}H_{i} -$$

Salt-isomerism type phototropic dyes

Night Blue

Victoria Blue R

$$\bigcap_{H} N - \bigcap_{H} \bigcap_{H} N - \bigcap_{H} \bigcap$$

Brilliant Milling Blue B Brilliant Blue F & R Ex. Eriocyanine A

$$SO_{3} - CH_{2} \\ CH_{2} \\ CH_{2} \\ N - CH_{3} \\ N \\ N(CH_{3})_{2}$$

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Methyl Blue

$$N_0SO_3 - \sqrt{} NH - \sqrt{} C = \sqrt{} NH - \sqrt{} SO_2N_3$$

Aniline Blue

Eriochrome Cyanine R

Methyl Violet 6B

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

Iodine Green

Aniline Blue

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Wool Violet 5 BN

$$\begin{array}{c|c} & & & & & & \\ & & & & & \\ & & & & & \\ & & & & & \\ & & & & & \\ & & & & \\ & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ &$$

Wool Violet 4 EM

Light Green SF Yellowish

Iodine Violet

Methyl Violet

$$CH_{ij}$$
 N C CH_{ij} N C

Crystal Violet

Ethyl Violet

$$(C_1H_3)_2N - C = - \dot{N}(C_2H_3)_2$$

$$N(C_3H_3)_2$$

Acid Green L Extra

Erioviridene B

Light Green SF

Victoria Green (Malachite Green)

$$C = N(CH_3)_2$$
 $N(CH_3)_2$

Red-Violet 5R

Brilliant Green "B"

Di-[4(N,N-diethylamine)phenyl]-[4-(N,N-diethylamine-2-methyl) phenyl] methyl carbonium

$$C_2H_5)_2N$$
 C
 $N(C_2H_5)_2$
 $N(C_2H_5)_2$

Tri-[4(N,N-dipropylamino)phenyl] methyl carbonium

$$C_{iHb}$$
 N C_{iHb} N C_{iHb} N C_{iHb} N N

Di-[4(N,N-diethylamino)phenyl]-[4(ethylamino)phenyl] methyl carbonium

$$\prod_{i} C_{i}H_{b}$$

Di-[4(N,N-diethylamino)phenyl]-[4(N,N-diethylamino)maphthyl] methyl carbonium

Di-[4(N,N-dimethylamino)phenyl]-[4(hydroxy)phenyl] methyl carbenium

Tri-[4(N-propylamino)phenyl] methyl carbonium

$$\bigcup_{\Pi} C_{J\Pi_{2}} N = \bigcup_{\Pi} C_{J\Pi_{2}} \sum_{\Pi} V = \bigcup_{\Pi} V = \bigcup_{\Pi}$$

Hectolene Blue DS-1398

Hectolene Blue DS-1823

Sevron Brilliant Red 4G

Di-[4(N,N-dimcthylamino)phenyi]-[4(hydroxy)phenyi]

methyl carbonium

Tri-[4(N-propylamino)phenyl] methyl carbonium

Hectolene Blue DS-1398

Hectolene Blue DS-1823

Sevron Brilliant Red 4G

Genacryl Red 6B

Genscryl Pink G Sevron Brilliant - Red B

Sevron Brilliant - Red 3B

1,5-bis-[4(N,N-dimethylamino)phenyl]-1,5-bis-

(phenyl)divinyl carbonium trifluoroscetate

$$(CH_1)_2N - (CH_2)_2 - (CH_2)_2$$

1,1,3,3-tetrakis[4(N,N-dimethylamino)phenyl] vinyl carbonium perchlorate

(CH₀)_N-

$$(CH_3)N$$
 $C=CH-C$
 $(CH_3)N$
 $N(CH_3)2$
 CO_4

1,5-bis-[4(N,N-dimethylamino)phenyl]-1,5-bis-(phenyl) divinyl carbonium p-toluenesulfonate

$$\begin{array}{c|c} \text{Polyment of Date} \\ \hline \\ (CH_{J})_{I}N \end{array} = \begin{array}{c|c} CH - CH = CH - C \\ \hline \\ (CH_{J})_{I}N \end{array} = \begin{array}{c|c} CH_{1}CH_{2}CH_{2}CH_{3$$

1,7-bis[4(N,N-dimethylamino)phenyl]-1,7-bis-(2,4-dichlorophenyl) trivinyl carbonium perchlorate

$$(CH_{i})_{N} = \bigcup_{C=CH=CH=CH=CH=CH=C} \bigcap_{C} \bigcap_{CH_{i}} \bigcap_{CH_{i}$$

ClO₄—

Di-[4(N,N-dimethylamino)phenyl vinyl]-[2,4-diphenyl-6-methane thiopyran] methyl carbonium perchlorate

ClO₄—

1,7-bis-[4(N,N-dimethylamino)phenyl]-1,7-bis-(4-chlorophenyl) trivinyl carbonium trifluoroacetate

1,1,3-tris-[4-(N,N-dimethylamino)phenyl] divinyl carbonium perchlorate

1,1,7,7-tetrakis-[4-(N,N-dimethylamino)phenyl]

trivinyl carbonium perchlorate

$$(CH_{3})N - (CH_{3})N - (CH$$

1,3-bis-[4-(N,N-dimethylamino)phenyl]-1,3-bis-(phenyl) vinyl carbonium perchlorate

1,1,5,5-tetrakis-[4-(N,N-dimethylamino)phenyl] divinyl carbonium perchlorate

$$\begin{pmatrix} (CH_3)_2N & & & \\ C=CH-CH=CH-C & & & \\ (CH_3)_2N & & & & \\ \end{pmatrix} N(CH_3)_2 \\ CO_4-CH=CH-C & & & \\ CO_4-CH=CH-C & & \\ CO_4-CH-C & & \\ CO_4-C$$

1,5-bis-[4-(N,N-dimethylamino)phenyl]-1,5-bis-(phenyl) divinyl carbonium perchlorate

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1,7-bis-[4-(N,N-dimethylamino)phenyl]-1,7-bis-(phenyl) trivinyl carbonium trifluoroacetate

$$(CH_{J})_{N} = CH - CH - CH - CH - CH - CH - CH$$

CF₃COO-

1(1,3,3-trimethyl indoline)-2-[4-(N,N-dimethylamino)phenyl] ethylene carbonium perchlorate

CIO₄—

1(1,3,3-trimethyl indoline) 4-[4-(N,N-dimethylamino)phenyl] butylene carbonium perchlorate

CIO₄—

1,1,3,3-tetrakis-[4(N,N-diethylamino)phenyl] vinyl carbonium perchlorate

$$(C_{3H_{0})_{2}N} - \bigcup_{C=CH-C} = N(C_{3H_{0})_{2}} N(C_{3H_{0})_{2}}$$

$$(C_{3H_{0})_{2}N} - \bigcup_{C} = CH-C$$

$$N(C_{3H_{0})_{2}N} - \bigcup_{C} N(C_{3H_{0})_{2}} N(C_{3H_{0})_{2}}$$

1,1-bis-[4-(N,N-diethylamino)phenyl]-3,3-bis-

[4-(N,N-dimethylamino)phenyl] vinyl carbonium perchlorate

1,1,5,5-tetrakis-[4-(N,N-diethylamino)phenyl] divinyl carbonium perchlorate

$$(C_2H_3)_2N - \\ C = CH - CH = CH - \\ (C_2H_3)_2N - \\ N(C_2H_3)_2$$

C104-

1,1-bis-[4-(N,N-dimethylamino)phenyl]-3-[4-(amino) phenyl]-3-methylvinyl carbonium perchlorate

$$(CH_{i})_{2}N - C = CH - C$$

$$(CH_{3})_{2}N - CH_{3} - CH_{4} - CIO_{4} - C$$

Tris-[1,1-bis-[4(N,N-dimethylamino)phenyl] ethylene] methyl carbonium perchlorate

$$(CH_{i,j})N \longrightarrow CH - C$$

$$(CH_{i,j})N \longrightarrow CH - C$$

$$(CH_{i,j})N \longrightarrow CH - C$$

$$CH -$$

C104-

Tris-[1,1-bis-[4-(N,N-diethylamino)phenyl] ethylene] methyl carbonium perchlorate

$$(C_{i}H_{0})_{2}N = \sum_{C=CH-C=CH-C} N_{i}C_{3}H_{0}_{0}$$

$$(C_{i}H_{0})_{2}N = \sum_{C=C} N_{i}C_{3}H_{0}_{0}$$

$$(C_{i}H_{0})_{3}N = \sum_{C=C} C_{i}C_{4}$$

$$(C_{i}H_{0})_{3}N = \sum_{C=C} C_{i}C_{4}$$

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$1,1,5\text{-tris-}[4\text{-}(N,N\text{-dimethylamino})\text{phenyl}] \ divinyl$

carbonium perchlorate

$$(CH_1)_2N - C = CH - CH = CH - CH = N(CH_3)_2$$

$$(CH_3)_2N - CO_4 - CH_3$$

N[4-(N,N-dimethylamino) cinnamylidene] auramine

$$(CH_{i})_{2}N \longrightarrow CH = CH - CH = OH - CH_{i}$$

$$(CH_{i})_{2}N \longrightarrow CH_{i}$$

$$(CH_{i})_{2}N \longrightarrow CH_{i}$$

$$(CH_{i})_{2}N \longrightarrow CH_{i}$$

1,1-bis-[4-(N,N-dimethylamino)phenyl-3,4-bis-(phenyl)]-3,4-diszo butene carbonium

1,1,5,5-tetrakis-[4-(N,N-dimethylamino)phenyl]-2,3-diazo pentene carbonium

N-(N',N'-dimethylamino cinnamytidene)-N,N-diphenyl ammonium

Azo Polymethines

Dyes of the general structural type

$$\sum_{\substack{C = N - (CH = CH)_n - C}} \stackrel{R}{\underset{D}{\longleftarrow}} \stackrel{N}{\underset{N}{\longleftarrow}} \stackrel{N}{\underset{N}{\underset{N}{\longleftarrow}} \stackrel{N}{\underset{N}{\longleftarrow}} \stackrel{N}{\underset{N}{\underset{N}{\longleftarrow}} \stackrel{N}{\underset{N}{\longrightarrow}} \stackrel{N}{\underset{N}{\longleftarrow}} \stackrel{N}{\underset{N}{\underset{N}{\longrightarrow}} \stackrel{N}{\underset{N}{\underset{N}{\longrightarrow}} \stackrel{N}{\underset{N}{\longleftarrow}} \stackrel{N}{\underset{N}{\longrightarrow}} \stackrel{N}{$$

$$(CH_{i})_{i}N = \begin{pmatrix} CH_{i} \\ CH_{i} \\ CH_{i} \end{pmatrix}_{i}N = \begin{pmatrix} CH_{i} \\ CH_{i} \\ CH_{i} \\ CH_{i} \end{pmatrix}_{i}N = \begin{pmatrix} CH_{i} \\ C$$

- 31. (Original) The method of synthesis of the compound of claim 10 wherein the C moiety is any molecule which exhibits bleaching behavior with the B moiety and has an increased therapeutic effect or therapeutic ratio as a consequence of its delivery as part of a prodrug.
- 32. (Original) The method of synthesis of the compound of claim 29 wherein the C moiety has a nucleophilic group that bonds to the B moiety.
- 33. (Original) The method of synthesis of the compound of claim 32 wherein the C moiety is derivatized to have a nucleophilic group that bonds to the B moiety.
- 34. (Original) The method of synthesis of the compound of claim 33 wherein the C moiety is

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derivatized by at least one of the nucleophilic groups comprising cinnamate, sulfite, phosphate, carboxylate, thiol, amide, alkoxide, or amine.

35. (Currently Amended) The method of synthesis of the compound of claim 10 wherein the C moiety is at least one of the group of

Captopril

HS — CH₂ O N COOK

Prostaglandin E₂

2,3-dichloro-α-methylbenzylamine

3'-deoxy-S-adenosyl-Lhomocysteine

Sinefungin

3,5-diiodo-4-hydroxybenzoic scid

6,6'-dithiobis (9-B-D-riboturanosylpurine)

γ-aminobutyric acid

H2NCH2CH2CH2COOH

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Gabaculine

N-(5'-phosphopyridoxy)-4-aminobutyric acid

4-amino-hex-5-enoic acid

Baclofen

Adenosine

3-hydroxy-3-methylglutarate

Campactin

But-3-ynoyl-CoA

Surami

L-3-iodotyrosine

L-3-iodo-a-methyltyrosine

Disodium cromoglycate

Adenosine 3',5'-cyclic monophosphate

D,L-B-(5-hydroxy-3indolyl)-α-hydrazinopropionic acid

D,L-α-hydrazino-αmethyidops

a-methyldona

5-(3,4-dihydroxycinnamoyl)salicylic acid

N-(phosphonacetyl)-L-aspartate

P-glycolohydroxamate

5-(p-sulfamylphenylazosalicylic acid

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Coformycin

Formycin B

Thioinosinate

Phosphonoformate

Phosphonoacetate

Ridavirin

Sotalol

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Cimetidine

Fuscaric acid

2-mercaptoethylamine

Mimosine

HSCH₂CH₂NH₃*

U-7130

Iproninzid

Trans-4-aminoocrotonic

acid

 $H_2NCH_2CH\!=\!CHCOOH$

NSD 1055

Nicotinic acid

Kynurenic acid

Lentysine Orotic acid Polyoxin D H2NCOCH2CHCHCHCHC Cephalosporin Penicillin

36. (Canceled)

- 37. (Original) The method of synthesis of the compound of claim 1 wherein the C moiety comprises at least one of the group of herbicides, fungicides, mitticides, nematocides, fumigants, growth regulators, repellants, defoliants, rodenticides, molluscicides, algicides, desicants, antehelmintics, and bactericides.
- 38. (Currently Amended) The method of synthesis of the compound of claim 37 wherein the C moiety is <u>a pesticide</u> one from the those given in Chemical Week Pesticides Register, Robert

P. Ovellette and John A. King, 1977, McGraw-Hill Book Company.

39. (Withdrawn) A method of synthesis of a chemical compound having the formula (A-B-C)_X-P-E_V

where the A is a chemiluminescent moiety,

B is an energy acceptor moiety, and

C is a biologically active mojety, and

P is a substrate

E is an enzyme and x and y are integers

comprising the steps of

forming a benzophenone,

forming a diaryl ethylene,

attaching a phthalimide moiety to at least one of the aryl groups of the ethylene to form a phthalimide-ethylene conjugate,

condensing two ethylene-phthalimide conjugates to form a phthalimide-pentadiene conjugate,

converting the phthalimide to the phthalhydrazide by reaction with hydrazine to form a carrier compound, and

reacting the carrier compound with a biologically active moiety to form a corresponding conjugate,

```
reacting A-B-C with a polymer to form (A-B-C)_X-P, and reacting E with (A-B-C)_X-P to form (A-B-C)_X-P-E_V.
```

- 40. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the compound provides controlled extra cellular release of the C moiety.
- 41. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the C moiety comprises at least one of drugs and proteins including enzymes and hormones.
- 42. (Withdrawn) The method of synthesis of the compound of claim 41 wherein the C moiety comprises at least one insulin, erythropoietin, interleuken 2, interferon, growth hormone, atrial natriuretic factor, tissue plasminogen activator, an anti-inflammatory drug, an antihypertensive drug, an inotropic drug, and a contraceptive drug.

- 43. (Withdrawn) The method of synthesis of the compound of claim 40 wherein extraacellular drug release occurs when the prodrug reacts with cellular free radicals via a mechanism involving chemiluminescence, photochromism, and intramolecular energy transfer.
- 44. (Withdrawn) The method of synthesis of the compound of claim 41 wherein the pharmaceutical agent is at least one of the group of antilipidemic drugs, anticholesterol drugs, contraceptive agents, anticoagulants, anti-inflamatory agents, immuno-suppressive drugs, antiarrhythmic agents, antineoplastic drugs, antihypertensive drugs, epinephrine blocking agents, cardiac inotropic drugs, antidepressant drugs, diuretics, antifungal agents, antibacterial drugs, anxiolytic agents, sedatives, muscle relaxants, anticonvulsants, agents for the treatment of ulcer disease, agents for the treatment of asthma and hypersensitivity reactions, antithroboembolic agents, agents for the treatment of muscular dystrophy, agents to effect a therapeutic abortion, agents for the treatment of anemia, agents to improve allograft survival, agents for the treatment of disorders of purine metabolism, agents for the treatment of ischemic heart disease, agents for the treatment of opiate withdrawal, agents which activate the effects of secondary messenger inositol triphosphate, agents to block spinal reflexes, and antiviral agents including a drug for the treatment of AIDS.
- 45. (Withdrawn) The method of synthesis of the compound of claim 43 wherein the C moiety is released by an oxidation reduction reaction with the target cell's electron carriers or by reaction with free radicals produced as a consequence of electron transport.
- 46. (Withdrawn) The method of synthesis of the compound of claim 43 wherein A represents a functionality which undergoes at least one of

an oxidation reduction reaction where electrons are transferred directly between A and the target cell's electron carriers, and

a reaction with free radicals of oxygen which are produced as a consequence of electron transport

such that an excited state is produced in A as a consequence of its participation in one of these reactions.

47. (Withdrawn) The method of synthesis of the compound of claim 46 wherein A undergoes intramolecular energy transfer from its own excited state to the B functionality which is an energy acceptor.

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- 48. (Withdrawn) The method of synthesis of the compound of claim 47 wherein upon receiving energy from A, B achieves an excited state which relaxes through heterolytic cleavage of the covalent bond of B with C where C is a drug moiety which is released into the environment.
- 49. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the chemiluminescent molecule comprises at least one of the group of

molecules undergoing reaction involving peroxides and oxygen free radicals,

molecules undergoing reaction involving oxidation or reduction, and

molecules undergoing both reaction with peroxides and oxygen free radicals followed by an oxidation or reduction reaction.

- 50. (Withdrawn) The method of synthesis of the compound of claim 49 wherein the chemilluminescent molecule comprises at least one of the group of luminol and its derivatives, lucigenin and its derivatives, Lophine and its derivatives, acridinium esters and acridans, tetraphenylpyrrole, phthalhydrazides, acyloins, biacridinium salts, vinylcarbonyls, vinylnitriles, tetrakis (dimethylamino) ethylene, acyloperoxides, indoles, tetracarbazoles and active oxalates.
- 51. (Withdrawn) The method of synthesis of the compound of claim 49 wherein the chemiluminescent molecule comprises at least one of the group of ruthenium chelates 2, 6-diaminopyrene, or cation radicals and molecules which follow a Chemically Initiated Electron Exchange Luminescence mechanism such as certain dioxetans and dioxetanones.
- 52. (Withdrawn) The method of synthesis of the compound of claim 49 wherein the chemiluminescent molecule comprises at least one of the group of dioxene derivatives and other compounds that form a dioxetan by reaction with superoxide and then produce efficient chemiluminescence by a CIEEL mechanism.
- 53. (Withdrawn) The method of synthesis of the compound of claim 49 wherein the chemiluminescent molecule comprises at least one of the group of

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2,6-diaminopyrene

Aminophthalhydrazide

TABLE 1

Representative Chemiluminescent Molecules

Name

Structure

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Dioxene

Imidazole derivatives

$$R_1 \longrightarrow \begin{pmatrix} N & & & \\ &$$

Sulfonyloxamides

Indole derivatives

Tetrakis(dialkylamino)ethylene

2,5,7,8-tetraoxabicyclo-[4.2.0.] octane

$$R_2$$
 O
 O
 R_1
 O
 O

Dioxetan

$$R_1$$
 R_2
 R_3

Lucigenin

TABLE 1 continued

Representative Chemiluminescent Molecules

Name	Structure
Lophine	
Actidinium esters	N CHIO X
Active oxalate	
Tris-2,2'-bipyridinedi- chlororuthenium (II)	(N N N N N N N N N N N N N N N N N N N
Dioxetanone	$CI \xrightarrow{CI} CI$ $O - O$ R_1 R_2
Dipheyl peroxide	°

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- 54. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the B moiety is a photochromic compound.
- 55. (Withdrawn) The method of synthesis of the compound of claim 54 wherein the photochromic compound comprises one which demonstrate photochromic behavior with electromagnetic radiation and bleaching agents.
- 56. (Withdrawn) The method of synthesis of the compound of claim 55 wherein the A functionality is chemiluminescent, and the B functionality is such that the photodissociative drug release spectrum of B overlaps the chemiluminescence spectrum of A.
- 57. (Withdrawn) The method of synthesis of the compound of claim 54 wherein the photochromic compound comprises a cationic dye.
- 58. (Withdrawn) The method of synthesis of the compound of claim 57 wherein the cationic dye comprises at least one of a di and triarylmethane dyes, triarylmethane lactones and cyclic ether dyes, cationic indoles, pyronines, phthaleins, oxazines, thiazines, acridines, phenazines, and anthocyanidins, and cationic polymethine dyes and azo and diazopolymethines, styryls, cyanines, hemicyanines, dialkylaminopolyenes, and other related dyes.
- (Withdrawn) The method of synthesis of the compound of claim 57 wherein the cationic dye comprises at least one of

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Dioxene

Imidazole derivatives

$$\underset{R_{4}-\text{\mathbb{Z}}}{\underset{R_{4}-\text{\mathbb{Z}}}{\bigwedge}}\underset{R_{3}}{\overset{R_{2}}{\bigwedge}}$$

Sulfonyloxamides

Indole derivatives

$$\bigcap_{N \in \mathbb{N}} R_i$$

Tetrakis(dialkylamino)ethylene

$$\begin{array}{c|c} & R & R & R \\ R-N & N-R \\ R-N & N-R \\ R & R & R \end{array}$$

2,5,7,8-tetraoxabicyclo-[4,2,0,] octane

$$O$$
 O
 R_2
 O
 O
 O

Dioxetan

Lucigenin

60. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the C moiety is any molecule which exhibits bleaching behavior with the B moiety and has an increased therapeutic effect or therapeutic ratio as a consequence of its delivery as part of a prodrug.

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- 61. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the C moiety has a nucleophilic group that bonds to the B moiety.
- 62. (Withdrawn) The method of synthesis of the compound of claim 61 wherein the C moiety is derivatized to have a nucleophilic group that bonds to the B moiety.
- 63. (Withdrawn) The method of synthesis of the compound of claim 62 wherein the C moiety is derivatized by at least one of the nucleophilic groups comprising cinnamate, sulfite, phosphate, carboxylate, thiol, amide, alkoxide, or amine.
- 64. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the C moiety is at least one of the group of

Captopril

Prostaglandin E2

2,3-dichloro-α-methylbenzylamine

3*-deoxy-S-ndenosyl-Lhomocysteine

Sinefungin

3,5-diiodo-4-hydroxybenzoic seid

γ-aminobutyric acid

H2NCH2CH2CH2COOH

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Gabaculine

N-(5'-phosphopyridoxy)-4-aminobutyric acid

4-amino-hex-5-enoic acid

$$\begin{array}{c} CH_2 = CHCHCH_2CH_2COOH \\ | \\ NH_2 \end{array}$$

Bactofen

Adenosine

3-hydroxy-3-methylglutarate

Campactin

But-3-ynoyl-CoA

Suramin

$$\begin{array}{c} \text{SO}_{r} \\ \text{NH} \\ \text{C} \\ \text{O}_{r} \\ \text{NH} \\ \text{C} \\ \text{C} \\ \text{O}_{r} \\ \text{NH} \\ \text{C} \\ \text{NH} \\ \text{NII} \\ \text{C} \\ \text{C} \\ \text{NH} \\ \text{C} \\ \text{NH} \\ \text{C} \\ \text{NH} \\ \text{C} \\ \text{NH} \\ \text{C} \\ \text{C} \\ \text{NH} \\ \text{C} \\ \text{NH} \\ \text{C} \\ \text{NH} \\ \text{C} \\$$

L-3-iodotyrosine

L-3-iodo-α-methyltyrosine

Disodium cromoglycate

N-(phosphonacetyl)-L-aspartate

P-glycolohydroxamate

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Coformycin

HO-H_IC O

Formycin B

Thioinosinate

Phosphonoformate

Phosphonoscetate

Ridavirin

Sotatol

Cimetidine

we full that
$$\begin{array}{c} \text{HIC} \\ \text{CH}_{1} \\ \text{CH}_{2} \\ \text{CH}_{3} \\ \text{CH}_{5} \\ \text$$

Fuscaric acid

HSCH2CH2NH3*

2-mercaptoethylamine

Mimosine

NH₂ I ÇH₂CHCOOH

U-7130

Iproniazid

Trans-4-aminoocrotonic acid

$$H_2NCH_2CH = CHCOOH$$

NSD 1055

Nicotinic acid

Kynurenic acid

Leasysine

NIII

N

OH

CITICTICHCOOH

OH

OH

Folyoxin D

Cephalosporin

Penicillin

- 65. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the A-B-C moieties are attached to P by a bond between P and at least one of A and B.
- 66. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the E moieties are attached to (A-B-C)_X-P by a bond between E and at least one of A, B, and P.
- 67. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the E moieties are enzymes that react with a desired substrate and form substances that cause the

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release of C from A-B-C.

- 68. (Withdrawn) The method of synthesis of the compound of claim 39 wherein the E moieties are enzymes that react with a desired substrate and form peroxide or free radicals that cause the release of C from A-B-C.
- 69. (Withdrawn) The method of synthesis of the compound of claim 67 wherein the E moiety, substrate, and C moiety are at least one of the group of

glucose oxidase, glucose, and insulin, and

xanthine oxidase, xanthine, and tissue plasminogen activator (TPA).

70. (Withdrawn)A method of synthesis of a chemical compound having the formula (A-B-C)_X-P

where the A is a chemiluminescent moiety,

B is an energy acceptor moiety, and

C is a biologically active moiety, and

P is a substrate and x is an integer

comprising the steps of

forming a benzophenone,

forming a diaryl ethylene,

attaching a phthalimide moiety to at least one of the aryl groups of the ethylene to form a phthalimide-ethylene conjugate,

condensing two ethylene-phthalimide conjugates to form a phthalimide-pentadiene conjugate,

converting the phthalimide to the phthalhydrazide by reaction with hydrazine to form a carrier compound, and

reacting the carrier compound with a strong base such as an alkali hydride and the biologically active moiety to form a corresponding conjugate,

reacting A-B-C with a polymer to form (A-B-C)_X-P.

- 71. (Original) The method of synthesis of the compound of claim 1 wherein one or more of the moieties can be modified to further candidate components by addition of functional groups.
- (Currently Amended) The method of synthesis of the compound of claim 71 wherein the groups comprise at least on one of alkyl, cycloalkyl, alkoxycarbonyl, cyano, carbamoyl,

heterocyclic rings containing C, O, N, S, sulfo, sulfamoyl, alkoxysulfonyl, phosphono, hydroxyl, halogen, alkoxy, alkylthiol, acyloxy, aryl, alkenyl, aliphatic, acyl, carboxyl, amino, cyanoalkoxy, alkenvlthio. cyanoalkoxycarbonyl, carboxvalkylcarboxamido, diazonium. carbamoylalkoxycarbonyl, alkoxy carbonylamino, cyanoalkylamino, alkoxycarbonylalkylamino, sulfoalkylamino, alkylsulfamoylaklylamino, oxido, hydroxy alkyl, carboxy alkylcarbonyloxy, cyanoalkyl, carboxyalkylthio, arylamino, heteroarylamino, alkoxycarbonyl, alkylcarbonyloxy. alkoxycarbonylalkoxy, carbamoylalkoxy, carbamovlalkyl carbonyloxy, cyanoalkoxy. sulfoalkoxy, nitro, alkoxyaryl, halogenaryl, amino aryl, alkylaminoaryl, tolyl, alkenylaryl, allylaryl, alkenvloxvarvl, allyloxyaryl, cvanoarvl. carbamovlaryl. carboxvaryl, alkoxycarbonylaryl, alkylcarbonyoxyaryl, sulfoaryl, alkoxysulfoaryl, sulfamoylaryl, and nitroaryl.

73. (Currently Amended) The method of synthesis of the compound of claim1 wherein the compound has the structure of general formula

$$\begin{array}{c|c} & & & & & & \\ & & & & & \\ & & & & & \\ & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ &$$

where R1, R2, and R3 are functional groups and L is a linker.

74. (Original) The method of synthesis of the compound of claim 73 wherein the functionality A is at least one of aminophthalhydrazide derivatives, sulfonyloxamides and active oxalates,

the functionality B is at least one of 1,1,5,5-tetrakisarylpentadiene and 1,1,5-trisarylpentadiene derivatives,

the functionality C is a drug molecule such as Foscarnate, or ddc;, and

R is a functional group, and

L is a linker such as an aliphatic chain between A and B.

75. (Original) The method of synthesis of the compound of claim 74 wherein the L functionality is between one 20 carbon atoms.

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76. (Currently Amended) The method of synthesis of the compound of claim 1 wherein B is a 1,1,5-trisarylpentadiene derivative and the compound has the formula

where R¹, R², and R³ are functional groups and L is a linker.

77. (Currently Amended) The method of synthesis of the compound of claim 1 wherein A is a sulfonyloxamide or active oxalate and the compound has the formula

where R¹ and R² are functional groups and L is a linker.

- 78. (Original) The method of synthesis of the compound of claim 1 wherein a luminol derivative is directly attached through one or more amino groups to the aryl groups of a photochromic dye.
- 79. (Currently Amended) The method of synthesis of the compound of claim 78 wherein C comprises the formula of at least one of

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Foscarnet and NH₂

ddc , and A-B comprises the formula of at least one of

YY99811-1

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6a

C₃₇H₃₈N₅O₂ • ClO₄ M.W. 684.20

MTLJ-1

80. (Currently Amended) The method of synthesis of the compound of claim 78 wherein the compound comprises the formula

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MTLJ-1-Foscarnet

- 81. (Original) The method of synthesis of the compound of claim 1 wherein the hydrolyzable group that protects phthalhydrazide is at least one of acetyl and t-butyloxycarbonyl.
- 82. (Original) The method of synthesis of the compound of claim 1 wherein the aminophthalimide-substituted precursors for the dye are prepared through amination of an aryl halide such as palladium-catalyzed amination of aryl halides.
- 83. (Original) The method of synthesis of the compound of claim 1 wherein halo-substituted aryl groups of a starting B moiety or an intermediate are coupled with the aminophthalimide by methods such as the aryl amination under palladium catalysis to form the aminophthalimide-substituted precursors for the dye.
- 84. (Original) The method of synthesis of the compound of claim 1 wherein halo-substituted aryl groups of a starting phthalimide or an intermediate are coupled with the amino-substituted dye by methods such as the aryl amination under palladium catalysis to form the aminophthalimide-substituted precursors for the dye.
- 85. (Currently Amended) The method of synthesis of the compound of claims 83-and 84 wherein amino-substituted aryl groups are obtained by the amination of the halo-substituted compounds with an imine such as benzophenoneimine.
- 86. (Original) The method of synthesis of the compound of claim 1 wherein the

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aminophthalimide-attached dye is formed by the condensation of two aminophthalimide-attached ethylene molecules by reaction with triethyl orthoformate and a strong acid such as perchloric acid in acetic anhydride or acetic acid.

- 87. (Original) The method of synthesis of the compound of claim 1 wherein during the step of converting the phthalimide moiety to the aminophthalhydrazide to obtain A-B, the B moiety is protected from reaction with hydrazine by reacting with base such as sodium hydroxide, sodium methoxide and amines.
- 88. (Original) The method of synthesis of the compound of claim 87 wherein the phthalimide-B conjugate with a protected B moiety is refluxed with hydrazine in a suitable solvent such as an alcoholic solvent in inert atmosphere and then treated with acid such as perchloric acid, tetrafluroboric acid to regenerate a corresponding unaltered B moiety of the A-B conjugate.
- 89. (Original) The method of synthesis of the compound of claim 88 wherein A-B is reacted with one nucleophilic species of C to form A-B-C.
- 90. (Original) The method of synthesis of the compound of claim 1 wherein A-B is formed by starting with B comprising halo-substituted dyes, such as 1,5-bis(p-bromophenyl)-1,5-bis(p-dimethylaminophenyl)-pentadienium perchlorate.
- 91. (Original) The method of synthesis of the compound of claim 90 wherein cationic dyes are protected by reacting with base such as alkoxide and then coupled with the aminophthalimide by amination of aryl halide such as the palladium-catalyzed amination of aryl halide to obtain the alkoxide-protecting aminophthalimide-substituted dyes.
- 92. (Original) The method of synthesis of the compound of claim 91 wherein the aminophthalimide-B conjugate with a protected B moiety is refluxed with hydrazine in a suitable solvent such as an alcoholic solvent to convert the amino-phthalimide moiety to the aminophthalhydrazide moiety and then treated with acid to generate A-B.
- 93. (Original) The method of synthesis of the compound of claim 1 wherein the B comprises a tetraarylpolymethine, the aminophthalhydrazide precursor is an aminophthalic acid diester and the conjugate to form A-B is amino-phthalimideluminol-tetraaryl-polymethine.

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94. (Currently Amended) The method of synthesis of the compound of claim 1 wherein halosubstituted diarylketone are formed by at least one of direct acylation of arene with halosubstituted benzoyl halide under ferric chloride catalysis according to the following representative scheme

acylation according to the following representative scheme

- 95. (Original) The method of synthesis of the compound of claim 1 wherein a halosubstituted diarylketone is converted to the corresponding halo-substituted diarylketene such as halo-substituted 1,1-diarylethene.
- 96. (Original) The method of synthesis of the compound of claim 95 wherein the halosubstituted diarylketene is coupled with a precursor of amino-phthalhydrazide such as aminophthalimide, aminophthalic acid diester, by aryl amination such as the palladium-catalyzed amination of aryl halides to form the aminophthalimide-substituted 1,1-diarylethene.
- 97. (Original) The method of synthesis of the compound of claim 96 wherein the ethene is condensed with an orthoester such as triethylorthoformate in a nonaqueous solvent such as acetic anhdydide, containing an acid catalyst such as perchloric acid, tetrafluoroboric acid, to form the aminophthalimide-substituted tetraarylpolymethine dyc.
- 98. (Original) The method of synthesis of the compound of claim 97 wherein the aminophthalimide moiety is converted to the aminophthalhydrazide to obtain A-B.

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- 99. (Original) The method of synthesis of the compound of claim 98 wherein the B moiety is a cationic dye that is first protected by reacting with an anion such as hydroxide, methoxide and amine and the phthalimide-B conjugate with a protected B moiety is refluxed with hydrazine in a suitable solvent such as an alcoholic solvent in inert atmosphere and then treated with acid such as perchloric acid, tetrafluroboric acid to regenerate a corresponding unaltered B moiety of the A-B conjugate.
- 100. (Original) The method of synthesis of the compound of claim 99 wherein A-B is reacted with one nucleophilic species of a C such as a drug 2',3'-dideoxycytidine, Foscarnet, acycloguanosine to form A-B-C comprising a prodrug.
- 101. (Original) The method of synthesis of the compound of claim 95 wherein two halosubstituted diarylketene precursor compounds are condensed with an orthoester such as triethylorthoformate in a nonaqueous solvent such as acetic anhydride containing acid catalyst such as perchloric acid, tetrafluoroboric acid to form the halo-substituted tetraarylpolymethine dyes such as 1,5-bis(p-bromophenyl)-1,5-bis(p-dimethylaminophenyl)-pentadienium perchlorate.
- 102. (Original) The method of synthesis of the compound of claim 101 wherein the B moiety is a cationic dye that is protected by reacting with an anion such as alkoxide and then coupled with the aminophthalimide by amination of aryl halide such as the palladium-catalyzed amination of aryl halide to obtain the alkoxide-protected aminophthalimide-substituted tetraarylpolymethine dve.
- 103. (Original) The method of synthesis of the compound of claim 103 wherein the alkoxide-protected aminophthalimide-substituted tetraarylpolymethine dye is refluxed with hydrazine in a suitable solvent such as an alcoholic solvent to convert the amino-phthalimide moiety to the aminophthalhydrazide moiety and then treated with acid to generate A-B comprising a luminol-tetraarylpolymethine compound.
- 104. (Currently Amended) The method of synthesis of the compound of claim 1 comprising the general steps given by following representative formula

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1c: R = OCH₃ 1d: R = O(CH₂)₃CH₃ 1e: R = (CH₂)₃CH₃

CH₃MgBr

1a: R = N(CH₃)₂ 1b: R = H 1c: R = OCH₃ 1d: R = O(CH₂)₃CH₃ 1e: R = (CH₂)₃CH₃ 2a: R = N(CH₃)₂ 2b: R = H 2c: R = OCH₃ 2d: R = O(CH₂)₃CH₃ 2e: R = (CH₂)₃CH₃

4-(*N*-ethylamino)-*N*-methylphthalimide
Pd(OAc)₂, P(*t*-Bu)₃, *t*-BuONa

-

3a: R = N(CH₃)₂ 3b: R = H 3c: R = OCH₃ 3d: R = O(CH₂)₃CH₃ 3e: R = (CH₂)₃CH₃

3-(N-Ethylamino)-N,6-dimethylphthalimide

5f

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3a: R = N(CH₃)₂

3b: R = H

3c: R = OCH₃

3d: $R = O(CH_2)_3CH_3$

3e: R = (CH₂)₃CH₃

1) KOH, 2) H₂NNH₂, 3) HX

4a: R = N(CH₃)₂, X = CIO₄ 4b: R = H, X = BF₄ 4c: R = OCH₃, X = BF₄ 4d: R = O(CH₂)₃CH₃, X = BF₄ 4e: R = (CH₂)₃CH₃, X = BF₄

5e: R = (CH₂)₃CH₃, X = BF₄

105. (Original) The method of synthesis of the compound of claim 1 wherein the A

functionality comprises a phthalhydrazide such as a luminol derivative and the B functionality comprises a photochromic dye wherein A is attached to aryl groups of B comprising the steps of

forming a diaryl ketone,

forming a diaryl ketene from the diaryl ketone,

forming a protected aminophthalhydrazide such as aminophthalimide or aminophthalic acid diester.

adding a hydrocarbon linker to the protected aminophthalhydrazide, and

attaching the protected aminophthalhydrazide through the molecular linker to the aryl groups of diarylketene to form the precursor aminophthalimide-linked diarylketene, and reacting according to at least one of

- (a) forming the A functionality from the precursor, and condensing two molecules of B precursor linked to A to form A-B, and
- (b) condensing two precursor aminophthalimide-linked diarylketene molecules to form A precursor linked to B, and

forming the A functionality from the A precursor to form A-B.

- 106. (Original) The method of synthesis of the compound of claim 105 wherein the diaryl ketone is formed by a classical Friedel-Crafts acylation between a benzoyl halide and aryl compound with a hydrocarbon linker having a leaving group.
- 107. (Original) The method of synthesis of the compound of claim 106 wherein the aryl compound with a hydrocarbon linker having a leaving group comprises at least one of a halogenated-alkyl-aryl ether and a halogenated-alkyl-aryl amine wherein the halogen is the leaving group.
- 108. (Original) The method of synthesis of the compound of claim 107 wherein the halogenated-alkyl-aryl ether comprises 2-bromoethoxybenzene to give an aryl ketone such as 4-(2-bromoethoxy)benzophenone.
- 109. (Original) The method of synthesis of the compound of claim 107 wherein the halogenated-aklyl-aryl amine comprises 2-bromoethyl aminobenzene to give an aryl ketone such as 4-(2-bromoethyl aminobenzophenone.
- 110. (Original) The method of synthesis of the compound of claim 105 wherein the diaryl ketone is converted to the corresponding diarylketene by reacting with a methylating reagent such

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as a methyl Grignard reagent, methyl lithium reagent, lithium dimethylcopper reagent and then dehydration with acid.

- 111. (Original) The method of synthesis of the compound of claim 110 wherein the diaryl ketone is converted to the corresponding diarylketene by reacting with methylmagnesium bromide and then dehydration with acid.
- 112. (Original) The method of synthesis of the compound of claim 105 wherein the diaryl ketone is converted to the corresponding diarylketene by a Wittig reaction.
- 113. (Original) The method of synthesis of the compound of claim 105 wherein a linker is attached to the protected aminophthalhydrazide by a reaction of a nucleophilic group of the linker or protected aminophthalhydrazide with a leaving group of the linker or protected aminophthalhydrazide.
- 114. (Original) The method of synthesis of the compound of claim 105 wherein a linker is attached to the protected aminophthalhydrazide by reaction to form a bond between at least one of a nitrogen, oxygen, or carbon atom of the linker and at least one of a nitrogen, oxygen, or carbon atom of group of the protected aminophthalhydrazide by an addition or a substitution reaction of a leaving group.
- 115. (Original) The method of synthesis of the compound of claim 114 wherein a linker is attached to the protected aminophthalhydrazide by a substitution reaction of at least one of a halogen, tosylate group, ester group with a nitrogen, oxygen, or carbon atom.
- 116. (Original) The method of synthesis of the compound of claim 105 wherein attaching the protected aminophthalhydrazide through the molecular linker to one of the aryl groups of diarylketene to form the precursor aminophthalimide-linked diarylketene is by a reaction of a nucleophilic group of the linker or aryl group of diarylketene with a leaving group of the linker or aryl group of diarylketene.
- 117. (Original) The method of synthesis of the compound of claim 116 wherein a linker is attached to the aryl group of diarylketene by reaction to form a bond between at least one of a nitrogen, oxygen, or carbon atom of the linker and at least one of a nitrogen, oxygen, or carbon atom of group of the protected aminophthalhydrazide by an addition or a substitution reaction of

a leaving group.

- 118. (Original) The method of synthesis of the compound of claim 117 wherein a linker is attached to the aryl group of diarylketene by a substitution reaction of at least one of a halogen, tosylate group, ester group with a nitrogen, oxygen, or carbon atom.
- 119. (Original) The method of synthesis of the compound of claim 105 wherein the precursor aminophthalimide-linked diarylketene is further reacted by condensation of two aminophthalimide-linked diarylketenes with an orthoester to form B linked to the A precursor.
- 120. (Original) The method of synthesis of the compound of claim 119 wherein condensing reagent is triethylorthoformate.
- 121. (Currently Amended) The method of synthesis of the compound of claim 119 wherein the precursor aminophthalimide-linked diarylketene comprises at least one of the formula

and the precursor of A-B comprises at least one of the formula

122. (Original) The method of synthesis of the compound of claim 119 wherein the

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phthalimide moiety of the A precursor is converted to the phthalhydrazide A functionality by treating with hydrazine, forming A-B.

- 123. (Original) The method of synthesis of the compound of claims 122 wherein the B functionality is protected by reacting with an anion such as hydroxide, methoxide and amine, the A-B precursor is refluxed with hydrazine in a suitable solvent such as an alcoholic solvent in inert atmosphere and then treated with acid such as perchloric acid, tetrafluroboric acid to form A-B.
- 124. (Original) The method of synthesis of the compound of claim 105 wherein the phthalimide moiety of the A precursor of the precursor aminophthalimide-linked diarylketene is converted to the phthalhydrazide A functionality by treating with hydrazine, forming A attached to a B precursor.
- 125. (Original) The method of synthesis of the compound of claim 124 wherein the A-linked diarylketene is further reacted by condensation of two A-linked diarylketenes with an orthoester to form A-B.
- 126. (Original) The method of synthesis of the compound of claim 125 wherein condensing reagent is triethylorthoformate.
- 127. (Currently Amended) The method of synthesis of the compound of claim 126 wherein the A-linked diarylketene comprises at least one of the formula

18a: R=OCH₃ 18b: R=O(CH₂)₂CH₂

18c: R=(CH₂)₃CH₃

18d: $R=N(CH_3)_2$

and A-B comprises at least one of the formula

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20a: R=OCH₃ 20b: R=O(CH₂)₃CH₃ 20c: R=(CH₂)₃CH₃ 20d: R=N(CH₃)₂

- 128. (Currently Amended) The method of synthesis of the compound <u>according to any one</u> of claims 123 and 125 further comprising the step of reacting the B functionality with one nucleophilic species of a C functionality such as Foscarnet to form A-B-C.
- 129. (Currently Amended) The method of synthesis of the compound of claim 105 comprising the general steps given by following representative formula

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$$\begin{array}{c} O \\ R \\ & +$$

130. (Currently Amended) The method of synthesis of the compound of claim 105

comprising the general steps given by following representative formula

13

3

12d

131. (Original) The method of synthesis of the compound of claim 1 wherein the A

functionality comprises a phthalhydrazide such as a luminol derivative and the B functionality comprises a triarylpolymethine photochromic dye wherein A is attached to aryl groups of B comprising the steps of

forming a diaryl ketone,

forming a diaryl ketene from the diaryl ketone.

forming a protected aminophthalhydrazide such as aminophthalimide or aminophthalic acid diester.

adding a hydrocarbon linker to the protected aminophthalhydrazide, and

attaching the protected aminophthalhydrazide through the molecular linker to the aryl groups of diarylketene to form the precursor aminophthalimide-linked diarylketene, and reacting according to at least one of

- (a) forming the A functionality from the precursor, and condensing the A-linked diarylketene with an aryl alkene aldehyde to form A-B, and
- (b) condensing the precursor aminophthalimide-linked diarylketene with an aryl alkene aldehyde to form A precursor linked to B, and

forming the A functionality from the A precursor to form A-B.

- 132. (Original) The method of synthesis of the compound of claim 131 wherein the diaryl ketone is formed by a classical Friedel-Crafts acylation between a benzoyl halide and aryl compound with a hydrocarbon linker having a leaving group.
- 133. (Original) The method of synthesis of the compound of claim 132 wherein the aryl compound with a hydrocarbon linker having a leaving group comprises at least one of a halogenated-alkyl-aryl ether and a halogenated-alkyl-aryl amine wherein the halogen is the leaving group.
- 134. (Original) The method of synthesis of the compound of claim 133 wherein the halogenated-alkyl-aryl ether comprises 2-bromoethoxybenzene to give an aryl ketone such as 4-(2-bromoethoxy)benzophenone.
- 135. (Original) The method of synthesis of the compound of claim 133 wherein the halogenated-aklyl-aryl amine comprises 2-bromoethyl aminobenzene to give an aryl ketone such as 4-(2-bromoethyl amino)benzophenone.
- 136. (Original) The method of synthesis of the compound of claim 131 wherein the diaryl

ketone is converted to the corresponding diarylketene by reacting with a methylating reagent such as a methyl Grignard reagent, methyl lithium reagent, lithium dimethylcopper reagent and then dehydration with acid.

- 137. (Original) The method of synthesis of the compound of claim 136 wherein the diaryl ketone is converted to the corresponding diarylketene by reacting with methylmagnesium bromide and then dehydration with acid.
- 138. (Original) The method of synthesis of the compound of claim 131 wherein the diaryl ketone is converted to the corresponding diarylketene by a Wittig reaction.
- 139. (Original) The method of synthesis of the compound of claim 131 wherein a linker is attached to the protected aminophthalhydrazide by a reaction of a nucleophilic group of the linker or protected aminophthalhydrazide with a leaving group of the linker or protected aminophthalhydrazide.
- 140. (Original) The method of synthesis of the compound of claim 131 wherein a linker is attached to the protected aminophthalhydrazide by reaction to form a bond between at least one of a nitrogen, oxygen, or carbon atom of the linker and at least one of a nitrogen, oxygen, or carbon atom of group of the protected aminophthalhydrazide by an addition or a substitution reaction of a leaving group.
- 141. (Original) The method of synthesis of the compound of claim 140 wherein a linker is attached to the protected aminophthalhydrazide by a substitution reaction of at least one of a halogen, tosylate group, ester group with a nitrogen, oxygen, or carbon atom.
- 142. (Original) The method of synthesis of the compound of claim 131 wherein attaching the protected aminophthalhydrazide through the molecular linker to one of the aryl groups of diarylketene to form the precursor aminophthalimide-linked diarylketene is by a reaction of a nucleophilic group of the linker or aryl group of diarylketene with a leaving group of the linker or aryl group of diarylketene.
- 143. (Original) The method of synthesis of the compound of claim 142 wherein a linker is attached to the aryl group of diarylketene by reaction to form a bond between at least one of a nitrogen, oxygen, or carbon atom of the linker and at least one of a nitrogen, oxygen, or carbon

atom of group of the protected aminophthalhydrazide by an addition or a substitution reaction of a leaving group.

- 144. (Original) The method of synthesis of the compound of claim 143 wherein a linker is attached to the aryl group of diarylketene by a substitution reaction of at least one of a halogen, tosylate group, ester group with a nitrogen, oxygen, or carbon atom.
- 145. (Original) The method of synthesis of the compound of claim 131 wherein the precursor aminophthalimide-linked diarylketene is further reacted by condensation with an aryl alkene aldehyde in a nonaqueous solvent, containing an acid catalyst to form B linked to the A precursor.
- 146. (Original) The method of synthesis of the compound of claim 145 wherein the precursor aminophthalimide-linked diarylketene is an aminophthalimide-substituted 1,1-diarylethene,

the aryl alkene aldehyde is a p-aminophenyl alkene aldehyde such as p-(dimethylamino)cinnamaldehyde,

the nonaqueous solvent is acetic anhydride,

the acid catalyst is at least one of perchloric acid and tetrafluoroboric acid, and

the B linked to the A precursor comprises a aminophthalimide-substituted multiarylpolymethine dye.

147. (Currently Amended) The method of synthesis of the compound of claim 145 wherein the precursor aminophthalimide-linked diarylketene comprises at least one of the formula

the aryl alkene aldehyde has the formula

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4-(Dimethylamino)cinnamaldehyde and

the precursor of A-B comprises at least one of the formula

- 148. (Original) The method of synthesis of the compound of claim 145 wherein the phthalimide moiety of the A precursor is converted to the phthalhydrazide A functionality by treating with hydrazine, forming A-B.
- 149. (Original) The method of synthesis of the compound of claims 148 wherein the B functionality is protected by reacting with an anion such as hydroxide, methoxide and amine, the A-B precursor is refluxed with hydrazine in a suitable solvent such as an alcoholic solvent in inert atmosphere and then treated with acid such as perchloric acid, tetrafluroboric acid to form A-B.
- 150. (Original) The method of synthesis of the compound of claim 131 wherein the phthalimide moiety of the A precursor of the precursor aminophthalimide-linked diarylketene is converted to the phthalhydrazide A functionality by treating with hydrazine, forming A attached to a B precursor.
- 151. (Original) The method of synthesis of the compound of claim 150 wherein the A-linked diarylketene is further reacted by condensation with an aryl alkene aldehyde in a nonaqueous

solvent, containing an acid catalyst to form A-B.

152. (Original) The method of synthesis of the compound of claim 151 wherein the A-linked diarylketene is an aminophthalhydrazide-substituted 1,1-diarylethene.

the aryl alkene aldehyde is a p-aminophenyl alkene aldehyde such as p-(dimethylamino)cinnamaldehyde,

the nonaqueous solvent is acetic anhydride,

the acid catalyst is at least one of perchloric acid and tetrafluoroboric acid, and A-B comprises a aminophthalhydrazide-substituted multiarylpolymethine dye.

153. (Currently Amended) The method of synthesis of the compound of claim 152 wherein the A-linked diarylketene comprises at least one of the formula

the aryl alkene aldehyde has the formula

4-(Dimethylamino)cinnamaldehyde, and

A-B comprises at least one of the formula

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- 154. (Currently Amended) The method of synthesis of the compound <u>according to any one</u> of claims 149 and 151 further comprising the step of reacting the B functionality with one nucleophilic species of a C functionality such as Foscarnet to form A-B-C.
- 155. (Currently Amended) The method of synthesis of the compound of claim 131 comprising the general steps given by following representative formula

3d: R = O(CH₂)₃CH₃ 3e: R = (CH₂)₃CH₃

23c: R = OCH₃

23d: $R = O(CH_2)_3CH_3$

23e: R = (CH₂)₃CH₃

24c: R = OCH₃

24d: R = O(CH₂)₃CH₃

24e: R = (CH₂)₃CH₃

24f

156. (Original) The method of synthesis of the compound of claim 131 comprising the general steps given by following representative formula

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